PCT

WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification 6:

C12N 15/00, 5/10, A01K 67/027, G01N 33/50, 33/68 // C12N 15/33, C07K 14/025, C12N 15/12, C07K 14/475, 14/495

(11) International Publication Number:

WO 97/39117

(43) International Publication Date:

23 October 1997 (23.10.97)

(21) International Application Number:

PCT/GB97/01063

(22) International Filing Date:

17 April 1997 (17.04.97)

(30) Priority Data:

9607953.8

17 April 1996 (17.04.96)

GB

(71) Applicant (for all designated States except US): THE UNI-VERSITY OF LIVERPOOL [GB/GB]; Senate House, Abercromby Square, Liverpool L69 3BX (GB).

(72) Inventors; and

(75) Inventors/Applicants (for US only): RUDLAND, Philip,
Spencer [GB/GB]; Dept. of Biological Sciences, The
University of Liverpool, Liverpool L69 3BX (GB). BARRACLOUGH, Barry, Roger [GB/GB]; Dept. of Biological
Sciences, The University of Liverpool, Liverpool L69 3BX
(GB). KILTY, Iain, Charles [GB/GB]; Dept. of Biological
Sciences, The University of Liverpool, Liverpool L69 3BX
(GB). DAVIES, Barry, Robert [GB/GB]; Addenbrooks
Hospital, P.O. Box 238, Cambridge CB2 2QQ (GB).
SCHMIDT, Guenter [DE/GB]; 13 Station Road, Cambridge
CB1 2JB (GB).

(74) Agent: W.P. THOMPSON & CO.; Coopers Building, Church Street, Liverpool L1 3AB (GB).

(81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, US, UZ, VN, ARIPO patent (GH, KE, LS, MW, SD, SZ, UG), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG).

Published

With international search report.



(54) Title: CONDITIONALLY IMMORTALISED CELL LINES DERIVED FROM TRANSGENIC ANIMALS

(57) Abstract

The invention relates to mammalian cell lines and transgenic mammals. More particularly, it relates to a method for producing a rat cell line, a method for producing a transgenic rat, a transgenic rat, a rat cell line, cells and tissue obtained therefrom and uses therefore. The cell line derived from a transgenic mammal comprises: (i) a conditional oncogene, transforming gene or immortalising gene or a cell cycle affecting gene; and (ii) a cell type specific promoter. They include a neuronal cell line in which the cell type specific promoter is an NF-L gene promoter, and a mammary cell line in which the cell type specific promoter is a MMTV gene promoter. The conditional oncogene, transforming gene or immortalising gene is preferably a SV40tsA58 gene.

onal Application No

PCT/GB 97/01063

A. CLASSIFICATION OF SUBJECT MATTER IPC 6 C12N15/00 C12N5/10 G01N33/68 G01N33/50 A01K67/027 //C12N15/33,C07K14/025,C12N15/12,C07K14/475,C07K14/495,

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 6 C12N A01K G01N C07K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 91 13150 A (LUDWIG INST CANCER RES ;MEDICAL RES COUNCIL (GB)) 5 September 1991	1-3,5,6, 10-17, 21-24
Y	see abstract	4,8,9, 19,20
	see page 1, line 1-13	
	see page 2, line 21 - page 3, line 9	
	see page 9, line 4-8	
	see page 12, line 1 - page 14, line 8 see page 15, line 14 - page 19, line 28	
	see page 22, line 28 - page 26, line 21	
	see page 29, line 1-6	1
	see page 29, line 26 - page 30, line 10	·
	see page 31, line 24 - page 34, line 20	
	see page 38, line 25 - page 40, line 10	
	-/	
	,	
		İ

* Special categories of cited documents: *A* document defining the general state of the art which is not considered to be of particular relevance	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
 *E' earlier document but published on or after the international filing date *L' document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) *O' document referring to an oral disclosure, use, exhibition or other means *P' document published prior to the international filing date but 	"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
later than the priority date claimed	*& document member of the same patent family Date of mailing of the international search report
Date of the actual completion of the international search 4 August 1997	0 8. 08. 97
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2	Authorized officer
NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Macchia, G

| X | Further documents are listed in the continuation of box C.

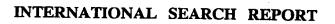
Patent family members are listed in annex.

r. onal Application No PCT/GB 97/01063

CIC	DOCUMENTS CONCIDED TO BUILD BY SVANT	
Category *	tion) DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X .	CELL, vol. 54, 1 September 1988, pages 105-115, XP002034399 MULLER ET AL.: "Single-step induction of mammary adenocarcinoma in transgenic mice bearing the activated c-neu oncogene" see page 105	1-3,5,7, 18
X	EP 0 169 672 A (HARVARD COLLEGE) 29 January 1986	1,2
Y A	see page 5, line 18-19 see page 6, line 1 - page 7, line 11 see page 17, line 7 - page 18, line 11 see page 19 - page 21; claims	19,20 3,7,18
Y	BIOCHEMICAL AND BIOPHYSICAL RESEARCH COMMUNICATIONS, vol. 192, no. 2, 30 April 1993, pages 465-470, XP002034398 REEBEN ET AL.: "Tissue-specific expression of rat light neurofilament promoter-driven reporter gene in transgenic mice" see abstract see page 467; figure 2 see page 470, line 6-8	4,8,9
Α	PROC. NATL. ACAD. SCI. USA, vol. 82, September 1985, pages 5880-5884, XP002034400 ROSS AND SOLTER: "Glucocorticoid regulation of mouse mammary tumor virus sequences in transgenic mice" cited in the application see page 5880, left-hand column see page 5883, right-hand column - page 5884	5,10
A	IN VIVO, vol. 8, no. 3, 1994, pages 263-270, XP002034877 MIZUNO M. ET AL.: "Normal and neoplastic mammary gland growth in MMTV-TGF alpha transgenic mice" see abstract	7,18
A	DATABASE WPI Section Ch, Week 9145 Derwent Publications Ltd., London, GB; Class B04, AN 91-329070 XP002034463 & JP 03 219 821 A (SNOW BRAND MILK PROD CO LTD), 27 September 1991 see abstract	12

PCT/GB 97/01063

tion) DOCUMENTS CONSIDERED TO BE RELEVANT		
Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No.
WO 95 11308 A (AMGEN INC) 27 April 1995 see page 33, line 22-28		13-16
WO 92 10563 A (UNIV PARIS VII) 25 June 1992		-1
WO 89 09816 A (MASSACHUSETTS INST TECHNOLOGY) 19 October 1989		
-		
	:	
	WO 95 11308 A (AMGEN INC) 27 April 1995 see page 33, line 22-28 WO 92 10563 A (UNIV PARIS VII) 25 June 1992 WO 89 09816 A (MASSACHUSETTS INST TECHNOLOGY) 19 October 1989	Citation of document, with indication, where appropriate, of the relevant passages WO 95 11308 A (AMGEN INC) 27 April 1995 see page 33, line 22-28 WO 92 10563 A (UNIV PARIS VII) 25 June 1992 WO 89 09816 A (MASSACHUSETTS INST TECHNOLOGY) 19 October 1989



.ternational application No.

PCT/GB 97/01063

Box i	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This Int	sternational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely: Remark: Although claim(s) 13-16 is(are) directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2.	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3.	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This Inte	ernational Searching Authority found multiple inventions in this international application, as follows:
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.	As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4.	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark o	The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

ed a donal Application No PCT/GB 97/01063

			
Patent document cited in search report	Publication date	Patent family member(s)	Publication date
WO 9113150 A	05-09-91	AU 676118 B AU 3305395 A AU 660604 B AU 7328691 A CA 2076345 A EP 0516664 A JP 5504066 T	27-02-97 11-01-96 06-07-95 18-09-91 21-08-91 09-12-92 01-07-93
EP 0169672 A	29-01-86	US 4736866 A DE 3586020 A JP 61081743 A US 5087571 A	12-04-88 17-06-92 25-04-86 11-02-92
WO 9511308 A	27-04-95	AU 7979194 A CA 2182958 A EP 0733118 A	08-05-95 27-04-95 25-09-96
WO 9210563 A	25-06-92	FR 2670215 A	12-06-92
WO 8909816 A	19-10-89	DE 68907220 T EP 0428519 A JP 3504799 T US 5338839 A US 5270191 A	27-01-94 29-05-91 24-10-91 16-08-94 14-12-93